

chain nodes :

12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33

ring nodes :

1 2 3 4 5 6 7 8 9 10 11

chain bonds :

1-11 2-20 3-19 4-12 6-13 7-22 7-33 8-14 8-23 9-15 9-24 11-21 12-17 12-18
14-25 15-16 25-26 25-31 26-27 26-30 26-32 27-28 27-29

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-11 8-9 9-10 10-11

exact/norm bonds :

1-2 1-6 1-11 2-3 3-4 4-5 4-12 5-6 6-13 7-8 7-11 7-33 8-9 8-14 9-10 10-11
14-25 25-31 26-30

exact bonds :

2-20 3-19 7-22 8-23 9-15 9-24 11-21 12-17 12-18 15-16 25-26 26-27 26-32 27-28
27-29

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:Atom
12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS 19:CLASS 20:CLASS
21:CLASS 22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS 27:CLASS 28:CLASS 29:CLASS
30:CLASS 31:CLASS 32:CLASS 33:CLASS

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10/607, 909

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	ENTRY	SESSION
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* The CA roles and document type information have been removed from *
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* available and contains the CA role and document type information. *
*

Structure search iteration limits have been increased. See HELP SLIMITS for details.

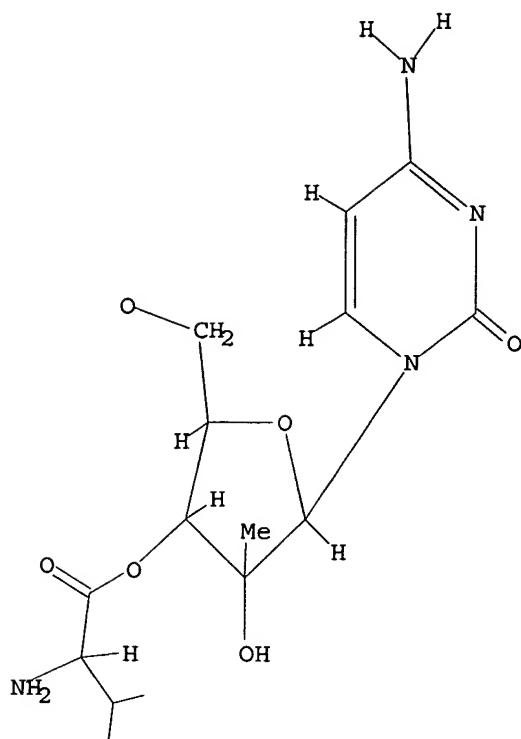
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L1 STRUCTURE UPLOADED

=> d 11
L1 HAS NO ANSWERS
L1 STR



Structure attributes must be viewed using STN Express query preparation.

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=> s 11 sss sam
SAMPLE SEARCH INITIATED 12:21:01 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 13 TO ITERATE
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```
100.0% PROCESSED 13 ITERATIONS 0 ANSWERS
SEARCH TIME: 00.00.01
```

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FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 44 TO 476
PROJECTED ANSWERS: 0 TO 0
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L2 0 SEA SSS SAM L1
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SEARCH TIME: 00.00.01
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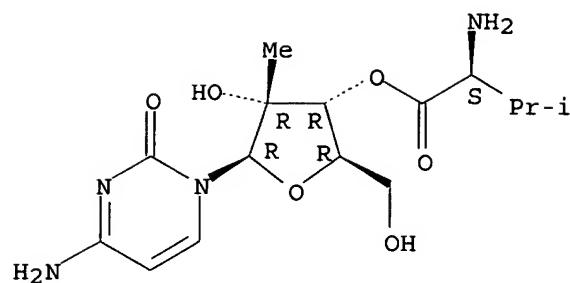
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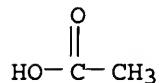
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L3 29 ANSWERS REGISTRY COPYRIGHT 2006 ACS on STN
IN L-Valine, 3'-ester with 2'-C-methylcytidine, acetate (salt) (9CI)
MF C15 H24 N4 O6 . x C2 H4 O2
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```
CM 1
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Absolute stereochemistry.



CM 2



HOW MANY MORE ANSWERS DO YOU WISH TO SCAN? (1):0

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COST IN U.S. DOLLARS	ENTRY	SESSION
FULL ESTIMATED COST	167.38	167.59

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=> s 13
 L4 8 L3

=> d bib abs hitstr 1-8 14

L4 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN
 AN 2006:149315 CAPLUS
 DN 144:205728
 TI Methods using a Type II interferon receptor agonist alone or in combination with a direct antiviral drug for treating hepatitis C virus infection
 IN Blatt, Lawrence M.
 PA Intermune, Inc., USA
 SO PCT Int. Appl., 139 pp.
 CODEN: PIXXD2
 DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2006016930	A2	20060216	WO 2005-US16927	20050513
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				

PRAI US 2004-571322P P 20040514

AB The invention provides methods for treating hepatitis C virus (HCV) infection; methods for reducing the incidence of complications associated with HCV and cirrhosis of the liver; and methods for reducing viral load, or reducing the time to viral clearance, or reducing morbidity or mortality in the clin. outcomes, in patients suffering from HCV infection. The methods generally involve administering to the individual a Type II interferon receptor agonist alone or in combination with a direct antiviral drug.

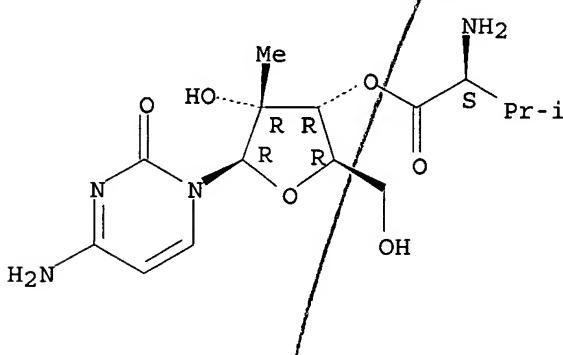
IT 640725-71-9, NM 283

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Usés)
(type II interferon receptor agonist alone or in combination with direct antiviral drug for treating hepatitis C virus infection)

RN 640725-71-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, dihydrochloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.



● 2 HCl

L4 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

AN 2005:684531 CAPLUS

DN 143:431740

TI Emerging drugs for chronic hepatitis C

AU Bhopale, Girish Mahadeorao; Nanda, Rabindra Kumar

CS Research and Development Division, Hindustan Antibiotics Limited, Pimpri, Pune, 411018, India

SO Hepatology Research (2005), 32(3), 146-153

CODEN: HPRSFH; ISSN: 1386-6346

PB Elsevier B.V.

DT Journal; General Review

LA English

AB A review. Hepatitis C virus (HCV) is a major cause of chronic hepatitis, liver cirrhosis and hepatocellular carcinoma worldwide. A combination therapy comprising pegylated interferon and ribavirin currently represents

the most effective therapy for chronic HCV infection. The limitations of this current therapy mainly its efficacy and significant side effects have prompted the development of new drugs. Few categories of therapeutic agents appear promising for future therapy, e.g. novel interferons, ribavirin analogs, antisense oligonucleotides, short interfering RNAs, ribozymes, enzyme inhibitors, immunomodulatory agents, antifibrotic agents, therapeutic vaccines and antibodies. Few drugs belong to afore-mentioned categories have already reached the different clin. phases of development. The present article highlights the status of current available therapies and emerging drugs for the treatment of hepatitis C.

IT 640725-71-9, NM 283

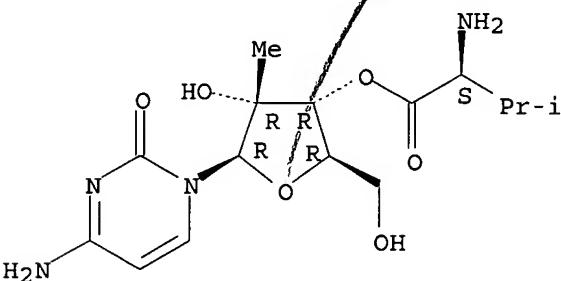
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(NM283 proved promising therapeutic effect in treating chronic hepatitis C patient)

RN 640725-71-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, dihydrochloride (9CI) (CA INDEX NAME)

Absolute stereochemistry:



● 2 HCl

RE.CNT 64 THERE ARE 64 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN
AN 2005:216597 CAPLUS

DN 142:291323

TI Compositions and methods for the treatment of severe acute respiratory syndrome (SARS)

IN Hardee, Greg; Dellamary, Luis

PA Isis Pharmaceuticals, Inc., USA

SO PCT Int. Appl., 217 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2005020885	A2	20050310	WO 2004-US16196	20040521
	WO 2005020885	A3	20050804		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,

GE, GH, GM, HF, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,

NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,

EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,

SN, TD, TG

PRAI US 2003-472774P P 20030521
AB The invention provides compns. and methods for treating a coronavirus

infection, especially a SARS CoV infection. The compns. comprise an antiviral nucleoside or mimetic thereof, or an antiviral antisense agent, in a form suitable for pulmonary or nasal delivery. The methods comprise administration to a patient in need thereof the effective amount of an antiviral composition by pulmonary or nasal instillation.

IT

640281-90-9

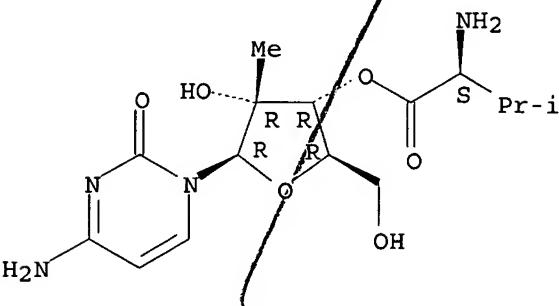
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(compns. and methods for treatment of severe acute respiratory syndrome)

RN 640281-90-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L4 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

AN 2004:515518 CAPLUS

DN 141:38814

TI Process for the production of 2'-branched nucleosides

IN Storer, Richard; Moussa, Adel; Chaudhuri, Narayan; Waligora, Frank

PA Idenix Cayman Limited, Cayman I.

SO PCT Int. Appl., 90 pp.

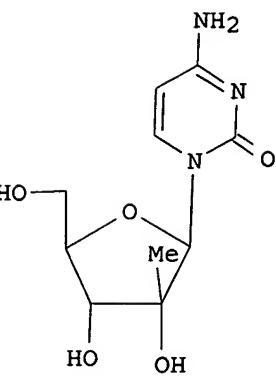
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 4

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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	WO 2004052899	A3	20050331		
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	RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	CA 2509687	AA	20040624	CA 2003-2509687	20031212
	US 2005020825	A1	20050127	US 2003-735408	20031212
	EP 1585529	A2	20051019	EP 2003-812993	20031212
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
	CN 1744903	A	20060308	CN 2003-80109576	20031212
	NO 2005003115	A	20050818	NO 2005-3115	20050624
PRAI	US 2002-432766P	P	20021212		
	US 2003-466194P	P	20030428		
	WO 2003-US39643	W	20031212		
OS	CASREACT				
GI	CASREACT				



AB The present invention provides an improved process for preparing ss-D and ss-L 2'-C-methyl-nucleosides and 2'-C-methyl-3'-O-ester nucleosides, e.g. I, via glycosylation of methylribonolactone with nucleobases.

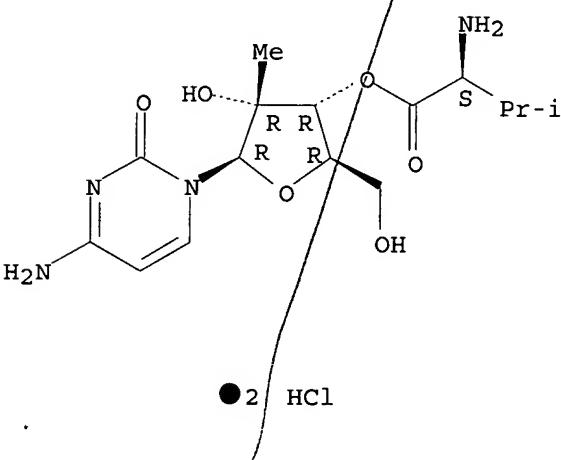
IT 640725-71-9P

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)
(process for production of 2'-branched nucleosides via glycosylation of methylribonolactone with nucleobases)

RN 640725-71-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, dihydrochloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L4 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

AN 2004:453348 CAPLUS

DN 141:17578

TI Treatment of Flaviviridae infection with 2'-branched nucleosides and another mutation-inducing drug such as interferon

IN Sommadossi, Jean-Pierre; La Colla, Paolo; Standring, David; Bichko, Vadim; Qu, Lin

PA Idenix (Cayman) Limited, Cayman I.; Universita Degli Studi Di Cagliari

SO PCT Int. Appl., 166 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2004046331	A2	20040603	WO 2003-US36714	20031117
	WO 2004046331	A3	20060302		

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TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
 RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
 BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
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 TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

CA 2506129 AA 20040603 CA 2003-2506129 20031117

US 2005031588 A1 20050210 US 2003-715729 20031117

EP 1576138 A2 20050921 EP 2003-796412 20031117

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK

BR 2003016363 A 20051004 BR 2003-16363 20031117

NO 2005002920 A 20050815 NO 2005-2920 20050615

PRAI US 2002-426675P P 20021115

WO 2003-US36714 W 20031117

OS MARPAT 141:17578

AB The present invention discloses a method for the treatment of Flaviviridae infection that includes the administration of a 2'-branched nucleoside, or a pharmaceutically acceptable prodrug and/or salt thereof, to a human in need of therapy in combination or alternation with a drug that directly or indirectly induces a mutation in the viral genome at a location other than a mutation of a nucleotide that results in a change from serine to a different amino acid in the highly conserved consensus sequence, XRXSGXXXT, of domain B of the RNA polymerase region, or is associated with such a mutation. The invention also includes a method to detect a mutant strain of Flaviviridae and a method for its treatment. Thus, in bovine viral diarrhea virus (BVDV)-infected MDBK cells treated with β -D-2'-methylcytidine, viruses resistant to the nucleoside appeared. The drug resistance was associated with a mutation in the NS5B gene which resulted in an S405T substitution in the encoded RNA-dependent RNA polymerase. These mutant viruses were sensitive to Intron A (interferon α -2b). Intron A and β -D-2'-methylcytidine exhibited synergistic inhibitory activity on BVDV growth in MDBK cells.

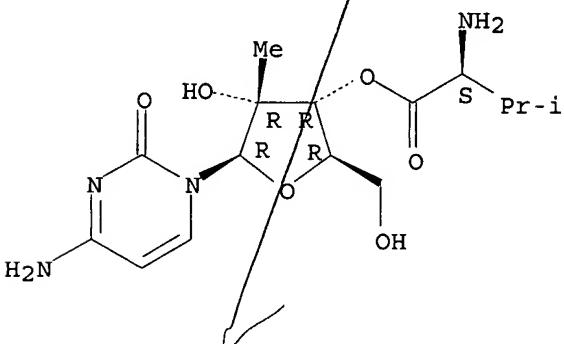
IT 640281-90-9

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (treatment of Flaviviridae infection with 2'-branched nucleosides and another mutation-inducing drug such as interferon)

RN 640281-90-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L4 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

AN 2004:20697 CAPLUS

DN 140:87662

TI 2'- and 3'-nucleoside prodrugs for treating Flaviviridae infections

IN Sommadossi, Jean-pierre; La Colla, Paolo; Storer, Richard; Gosselin, Gilles

PA Idenix (Cayman) Limited, Cayman I.; Centre National de la Recherche Scientifique; Universita Degli Studi di Cagliari

SO PCT Int. Appl., 2498 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 4

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI WO 2004003000 A2 20040108 WO 2003-IB3901 20030627

WO 2004003000 A3 20041104

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

CA 2490200 AA 20040108 CA 2003-2490200 20030627

AU 2003263412 A1 20040119 AU 2003-263412 20030627

EP 1525209 A2 20050427 EP 2003-761749 20030627

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK

JP 2005537242 T2 20051208 JP 2004-517162 20030627

CN 1761677 A 20060419 CN 2003-820501 20030627

WO 2005020884 A2 20050310 WO 2004-US15395 20040514

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

NO 2005000466 A 20050323 NO 2005-466 20050127

PRAI US 2002-392350P P 20020628

US 2002-392351P P 20020628

US 2003-466194P P 20030428

US 2003-470949P P 20030514

WO 2003-IB3901 W 20030627

OS MARPAT 140:87662

AB 2' And 3'-Prodrugs of 1', 2', 3', or 4'-branched β -D or β -L nucleosides, or their pharmaceutically acceptable salts and derivs., are described which are useful in the prevention and treatment of Flaviviridae infections and other related conditions. These modified nucleosides provide superior results against flaviviruses and pestiviruses, including hepatitis C virus and viruses generally that replicate through an RNA-dependent RNA reverse transcriptase. Compds., compns., methods and uses are provided for the treatment of Flaviviridae infection, including HCV infection, that include the administration of an effective amount of the prodrugs of the invention, or their pharmaceutically acceptable salts or derivs. These drugs may optionally be administered in combination or alternation with further antiviral agents to prevent or treat Flaviviridae infections and other related conditions. Preparation of compds. of the invention is included.

IT 640725-71-9P

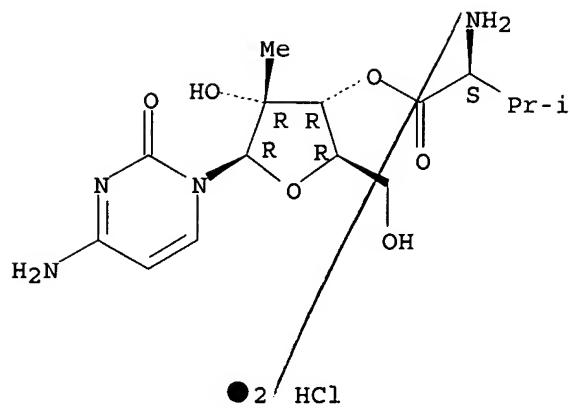
RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(nucleoside prodrugs for treating Flaviviridae infections)

RN 640725-71-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, dihydrochloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L4 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN

AN 2004:20696 CAPLUS

DN 140:77365

TI Preparation of modified 2'- and 3'-nucleoside prodrugs for treating Flaviviridae infections

IN Sommadossi, Jean-pierre; La Colla, Paolo; Storer, Richard; Gosselin, Gilles

PA Idenix (Cayman) Limited, Cayman I.; Universita degli studi di Cagliari; Centre National de la Recherche Scientifique

SO PCT Int. Appl., 201 pp.

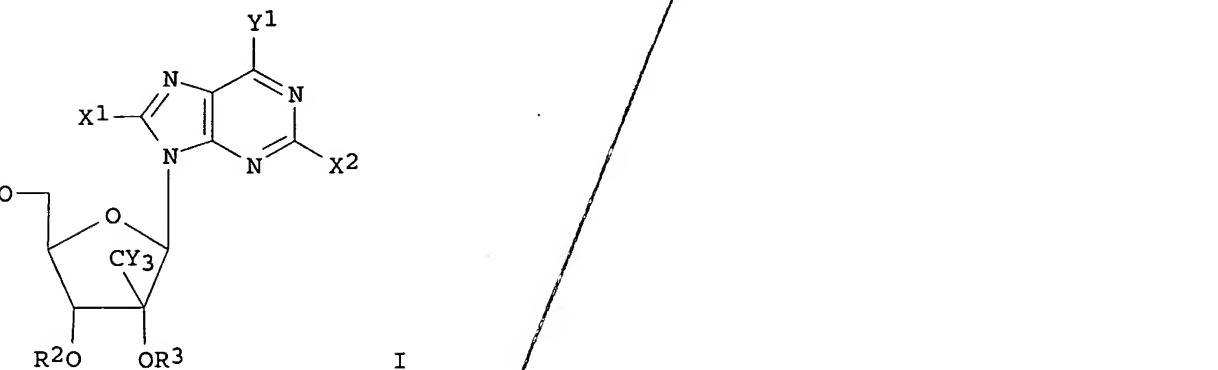
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 4

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2004002999	A2	20040108	WO 2003-IB3246	20030627
	WO 2004002999	A3	20040812		
	WO 2004002999	C1	20050217		
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	CA 2490191	AA	20040108	CA 2003-2490191	20030627
	AU 2003247084	A1	20040119	AU 2003-247084	20030627
	EP 1523489	A2	20050420	EP 2003-761744	20030627
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	JP 2005533817	T2	20051110	JP 2004-517158	20030627
	CN 1761677	A	20060419	CN 2003-820501	20030627
	WO 2005020884	A2	20050310	WO 2004-US15395	20040514
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
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PRAI	NO 2005000465	A	20050127	NO 2005-465	20050127
	US 2002-392350P	P	20020628		
	US 2002-392351P	P	20020628		
	US 2003-466194P	P	20030428		



AB 2' And/or 3' prodrugs of 1', 2', 3' or 4'-branched-nucleosides I, wherein R1-R3 are independently H, phosphate, alkyl, acyl, CO-alkyl, CO-aryl, CO-alkoxyalkyl, CO-aryloxyalkyl, CO-substituted aryl, sulfonate ester, benzyl, wherein the Ph group is optionally substituted with one or more substituents, alkylsulfonyl, arylsulfonyl, aralkylsulfonyl, lipid, amino acid, carbohydrate, peptide, cholesterol; Y1 is hydrogen, bromo, chloro, fluoro, iodo, CN, OH, OR4, NH2, NHR4, NR4R5, SH or SR4; X1 and X2 are independently alkyl, CH3, CF3, CY3, 2-Br-Et, CH2F, CH2Cl, CH2CF3, CF2CF3, CY2CY3, CH2OH, alkenyl, alkynyl, COOH, COOR4, COO-alkyl, COO-aryl, CO-O-alkoxyalkyl, CONH2, CONHR4, CON(R4)2, halo, CN, N3, OH, OR4, NH2, NHR4, NR4R5, SH or SR5; Y is independently H, halo; and each R4 and R5 is independently hydrogen, acyl, alkyl, lower alkyl, alkenyl, alkynyl or cycloalkyl, and their pharmaceutically acceptable salts and derivs. are described. These prodrugs are useful in the prevention and treatment of Flaviviridae infections, including HCV infection, and other related conditions. Compds. and compns. of the prodrugs of the present invention are described. Methods and uses are also provided that include the administration of an effective amount of the prodrugs of the present invention, or their pharmaceutically acceptable salts or derivs. These drugs may optionally be administered in combination or alteration with further anti-viral agents to prevent or treat Flaviviridae infections and other related conditions. Thus, antiviral activity of β -D-2'-C-methyl-7-methyl-6-phenyl-3,3a,5,8a-tetrahydro-1,3,4,5,7a-penta-aza-s-indacen-8-one is reported.

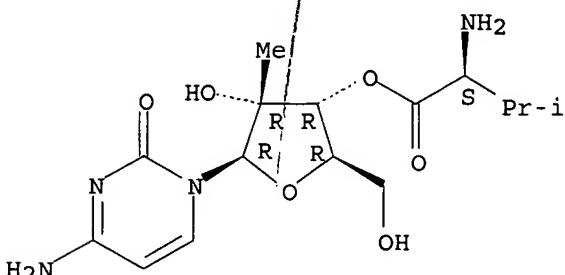
IT 640281-90-9P

RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of modified nucleoside prodrugs for treating flaviviridae infections)

RN 640281-90-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L4 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2006 ACS on STN
AN 2004:20443 CAPLUS
DN 140:70984

TI 2'-C-methyl-3'-O-L-valine ester ribofuranosyl cytidine for treatment of
flaviviridae infections

IN Sommadossi, Jean-Pierre; La Colla, Paolo
PA Idenix (Cayman) Limited, Cayman I.; Universita Degli Studi di Cagliari
SO PCT Int. Appl., 110 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 4

NY App

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2004002422	A2	20040108	WO 2003-US20431	20030627
	WO 2004002422	A3	20050407		

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LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PG,
PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR,
TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

CA 2489552 AA 20040108 CA 2003-2489552 20030627
US 2004077587 A1 20040422 US 2003-607909 20030627
EP 1536804 A2 20050608 EP 2003-762183 20030627

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IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK

JP 2005533824 T2 20051110 JP 2004-518041 20030627
WO 2005020884 A2 20050310 WO 2004-US15395 20040514

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
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NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
SN, TD, TG

NO 2005000490 A 20050127 NO 2005-490 20050127
PRAI US 2002-392351P P 20020628
US 2003-466194P P 20030428
US 2003-470949P P 20030514
WO 2003-US20431 W 20030627

OS MARPAT 140:70984

AB The 3'-L-valine ester of β -D-2'-C-methyl-ribofuranosyl cytidine provides superior results against flaviviruses and pestiviruses, including hepatitis C virus. Based on this discovery, compds., compns., methods and uses are provided for the treatment of flaviviridae, including HCV, that include the administration of an effective amount of val-mCyd or its salt, ester, prodrug or derivative, optionally in a pharmaceutically acceptable carrier. In an alternative embodiment, val-mCyd is used to treat any virus that replicates through an RNA-dependent RNA polymerase. Several examples are provided of the pharmacol., mechanism of action, metabolism, side effects, and clin. efficacy of the title compound

IT 640281-90-9D, salts 642075-50-1 642075-51-2

642075-52-3 642075-53-4 642075-54-5

642075-55-6 642075-56-7 642075-57-8

642075-58-9 642075-59-0 642075-60-3

642075-61-4 642075-62-5 642075-63-6

642075-64-7 642075-65-8 642075-66-9

642075-67-0 642075-68-1 642075-69-2

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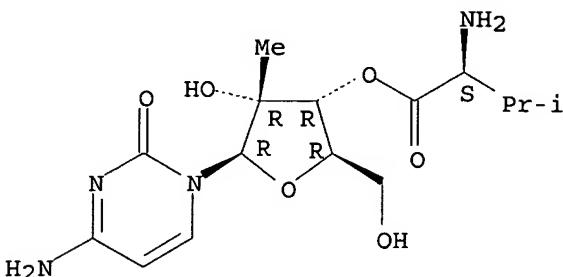
642075-77-2

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(ribofuranosylcytidine methylvaline ester combined with other
antivirals for treatment of flaviviridae infections)

RN 640281-90-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 642075-50-1 CAPLUS

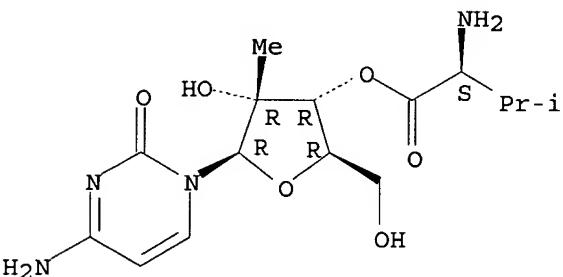
CN L-Valine, 3'-ester with 2'-C-methylcytidine, 4-methylbenzenesulfonate
(salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9

CMF C15 H24 N4 O6

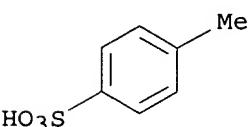
Absolute stereochemistry.



CM 2

CRN 104-15-4

CMF C7 H8 O3 S



RN 642075-51-2 CAPLUS

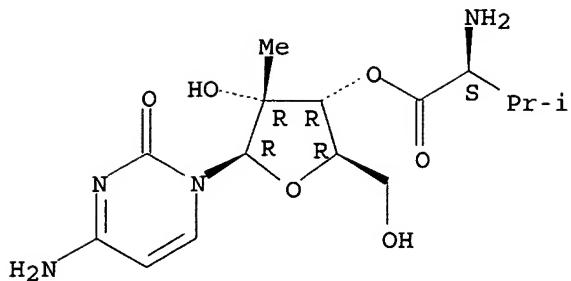
CN L-Valine, 3'-ester with 2'-C-methylcytidine, methanesulfonate (salt) (9CI)
(CA INDEX NAME)

CM 1

CRN 640281-90-9

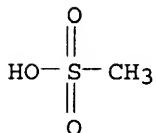
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 75-75-2
 CMF C H4 O3 S

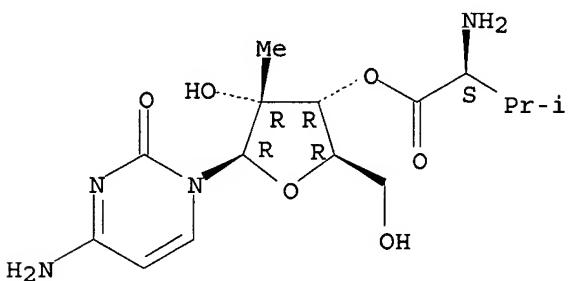


RN 642075-52-3 CAPLUS
 CN L-Valine, 3'-ester with 2'-C-methylcytidine, acetate (salt) (9CI) (CA INDEX NAME)

CM 1

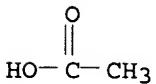
CRN 640281-90-9
 CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 64-19-7
 CMF C2 H4 O2



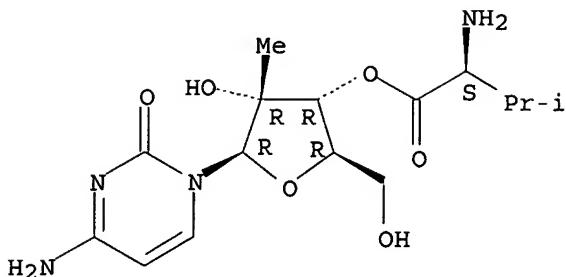
RN 642075-53-4 CAPLUS
 CN L-Valine, 3'-ester with 2'-C-methylcytidine, 2-hydroxy-1,2,3-propanetricarboxylate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9

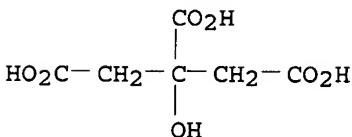
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 77-92-9
CMF C6 H8 O7



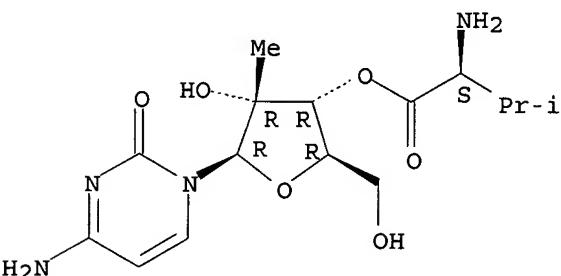
RN 642075-54-5 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, propanedioate (salt) (9CI)
(CA INDEX NAME)

CM 1

CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 141-82-2
CMF C3 H4 O4

$\text{HO}_2\text{C}-\text{CH}_2-\text{CO}_2\text{H}$

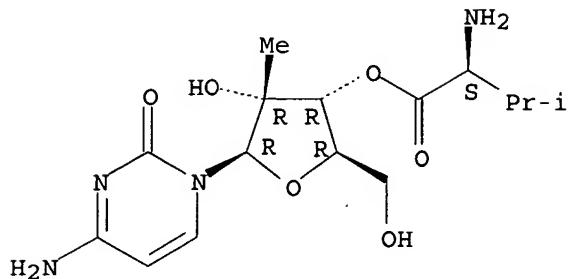
RN 642075-55-6 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, (2R,3R)-2,3-dihydroxybutanedioate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9
CMF C15 H24 N4 O6

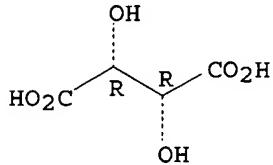
Absolute stereochemistry.



CM 2

CRN 87-69-4
CMF C4 H6 O6

Absolute stereochemistry.



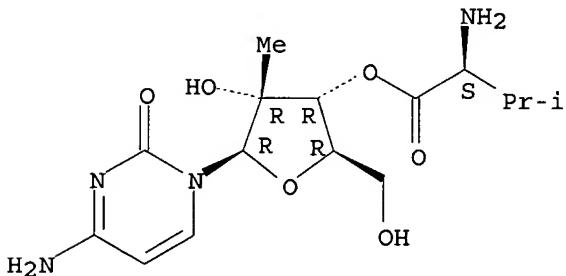
RN 642075-56-7 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, butanedioate (salt) (9CI)
(CA INDEX NAME)

CM 1

CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 110-15-6
CMF C4 H6 O4

HO₂C—CH₂—CH₂—CO₂H

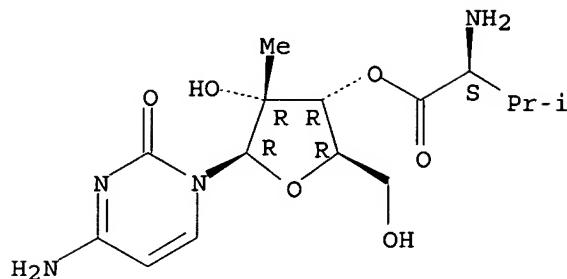
RN 642075-57-8 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, benzoate (salt) (9CI) (CA INDEX NAME)

CM 1

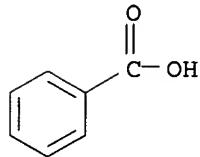
CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 65-85-0
CMF C7 H6 O2



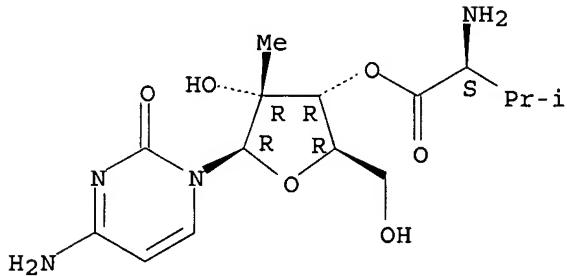
RN 642075-58-9 CAPLUS

CN L-Ascorbic acid, compd. with L-valine 3'-ester with 2'-C-methylcytidine
(9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9
CMF C15 H24 N4 O6

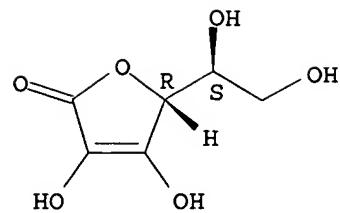
Absolute stereochemistry.



CM 2

CRN 50-81-7
CMF C6 H8 O6

Absolute stereochemistry.



RN 642075-59-0 CAPLUS

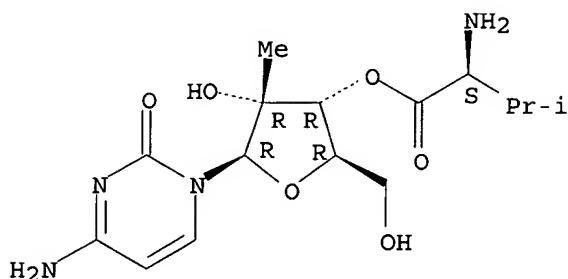
CN L-Valine, 3'-ester with 2'-C-methylcytidine, 2-oxopentanedioate (salt)
(9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9

CMF C15 H24 N4 O6

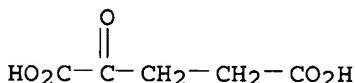
Absolute stereochemistry.



CM 2

CRN 328-50-7

CMF C5 H6 O5



RN 642075-60-3 CAPLUS

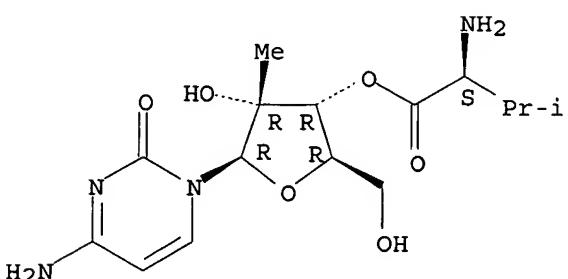
CN L-Valine, 3'-ester with 2'-C-methylcytidine, 2,3-dihydroxypropyl phosphate
(salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9

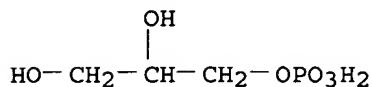
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 57-03-4
CMF C3 H9 06 P



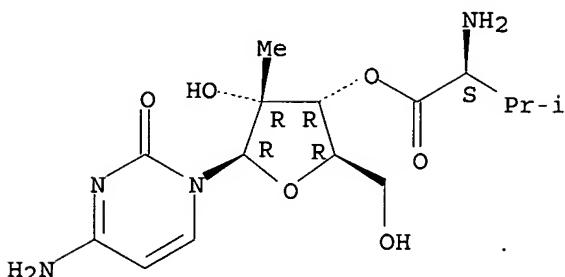
RN 642075-61-4 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, formate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9
CMF C15 H24 N4 06

Absolute stereochemistry.



CM 2

CRN 64-18-6
CMF C H2 O2

$$\text{O}=\text{CH}-\text{OH}$$

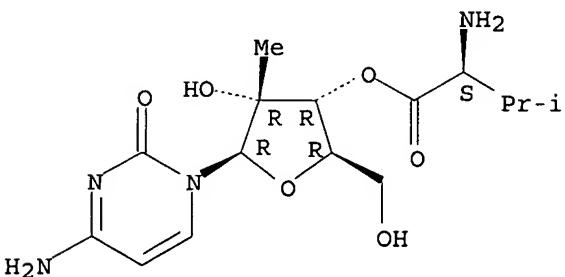
RN 642075-62-5 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, (2E)-2-butenedioate (salt)
(9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9
CMF C15 H24 N4 06

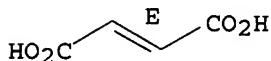
Absolute stereochemistry.



CM 2

CRN 110-17-8
CMF C4 H4 O4

Double bond geometry as shown.



RN 642075-63-6 CAPLUS

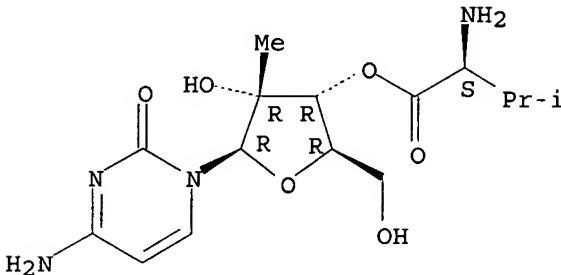
CN L-Valine, 3'-ester with 2'-C-methylcytidine, propanoate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9

CMF C15 H24 N4 O6

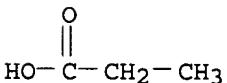
Absolute stereochemistry.



CM 2

CRN 79-09-4

CMF C3 H6 O2



RN 642075-64-7 CAPLUS

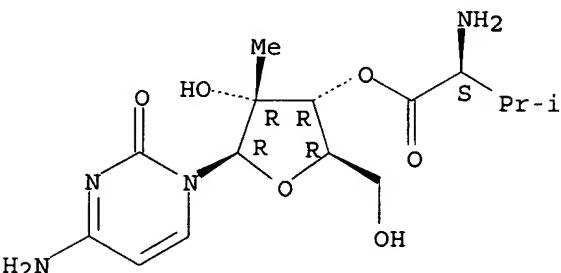
CN L-Valine, 3'-ester with 2'-C-methylcytidine, hydroxyacetate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9

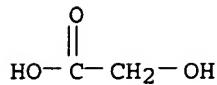
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 79-14-1
CMF C2 H4 O3



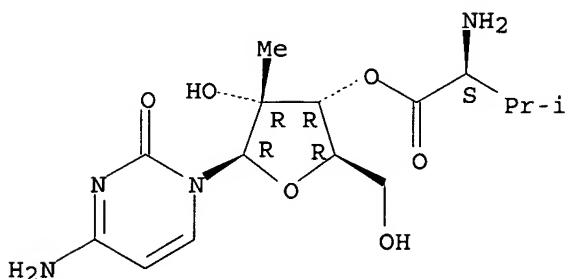
RN 642075-65-8 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, 2-hydroxypropanoate (salt)
(9CI) (CA INDEX NAME)

CM 1

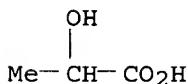
CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 50-21-5
CMF C3 H6 O3



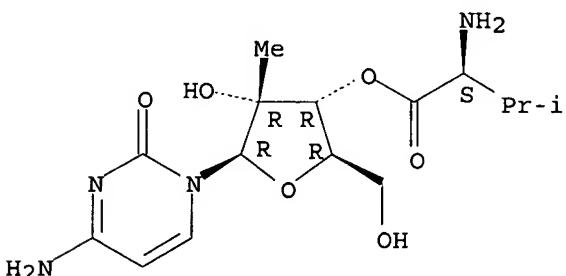
RN 642075-66-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, 2-oxopropanoate (salt) (9CI)
(CA INDEX NAME)

CM 1

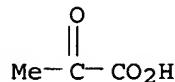
CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 127-17-3
CMF C3 H4 O3



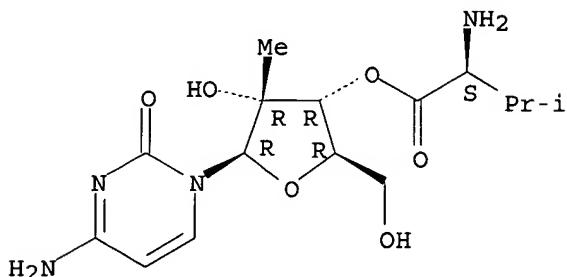
RN 642075-67-0 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, ethanedioate (salt) (9CI)
(CA INDEX NAME)

CM 1

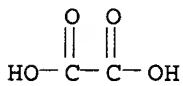
CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 144-62-7
CMF C2 H2 O4



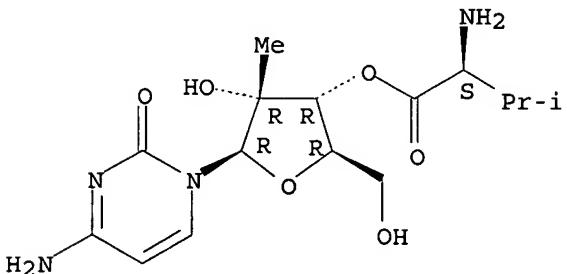
RN 642075-68-1 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, (2Z)-2-butenedioate (salt)
(9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9
CMF C15 H24 N4 O6

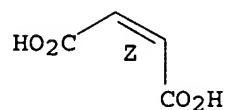
Absolute stereochemistry.



CM 2

CRN 110-16-7
CMF C4 H4 O4

Double bond geometry as shown.

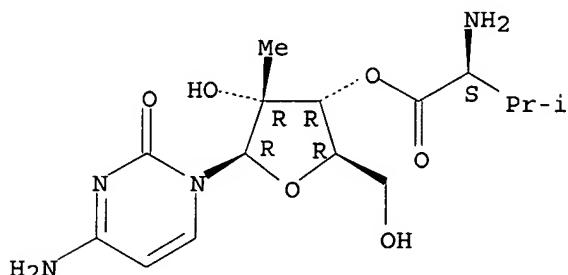


RN 642075-69-2 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, 2-hydroxybenzoate (salt)
(9CI) (CA INDEX NAME)

CM 1

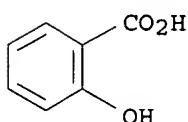
CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 69-72-7
CMF C7 H6 O3

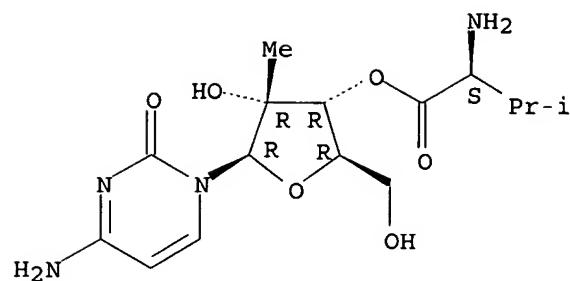


RN 642075-70-5 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, sulfate (salt) (9CI) (CA INDEX NAME)

CM 1

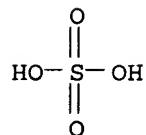
CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 7664-93-9
CMF H2 O4 S

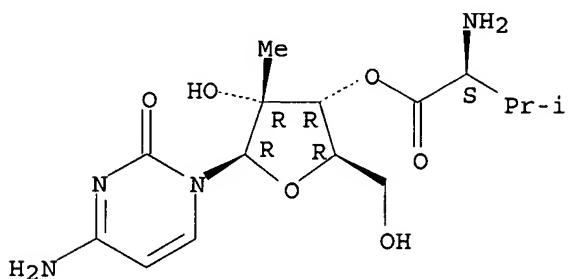


RN 642075-71-6 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, nitrate (salt) (9CI) (CA INDEX NAME)

CM 1

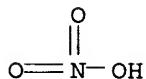
CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



CM 2

CRN 7697-37-2
CMF H N O3

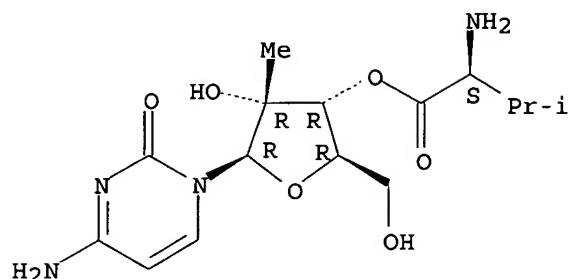


RN 642075-72-7 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, carbonate (salt) (9CI) (CA INDEX NAME)

CM 1

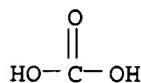
CRN 640281-90-9
CMF C15 H24 N4 O6

Absolute stereochemistry.



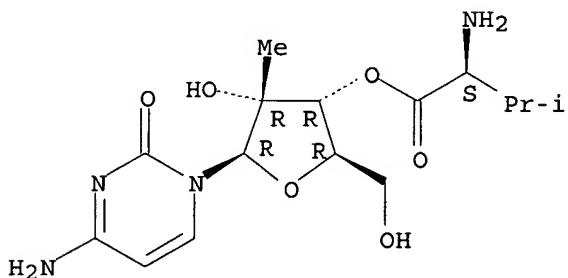
CM 2.

CRN 463-79-6
CMF C H2 O3



RN 642075-74-9 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, hydrobromide (9CI) (CA INDEX NAME)

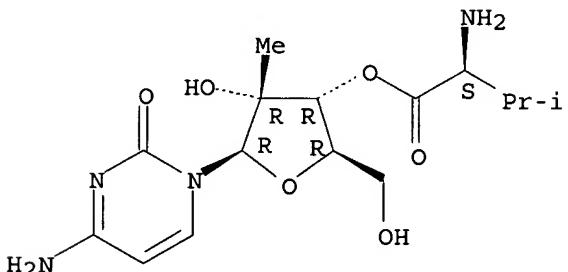
Absolute stereochemistry.



●x HBr

RN 642075-75-0 CAPLUS
CN L-Valine, 3'-ester with 2'-C-methylcytidine, hydriodide (9CI) (CA INDEX NAME)

Absolute stereochemistry.



●x HI

RN 642075-76-1 CAPLUS

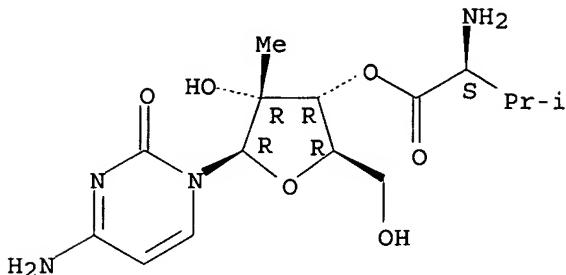
CN L-Valine, 3'-ester with 2'-C-methylcytidine, carbonate (2:1) (salt) (9CI)
(CA INDEX NAME)

CM 1

CRN 640281-90-9

CMF C15 H24 N4 O6

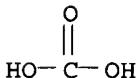
Absolute stereochemistry.



CM 2

CRN 463-79-6

CMF C H2 O3



RN 642075-77-2 CAPLUS

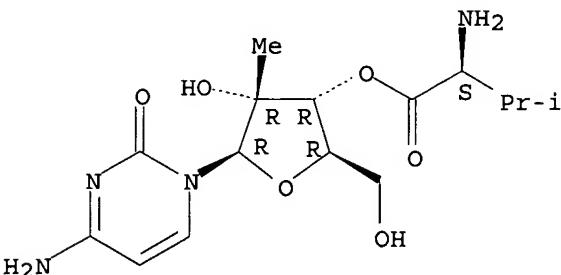
CN L-Valine, 3'-ester with 2'-C-methylcytidine, phosphate (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 640281-90-9

CMF C15 H24 N4 O6

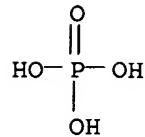
Absolute stereochemistry.



CM 2

CRN 7664-38-2

CMF H3 O4 P



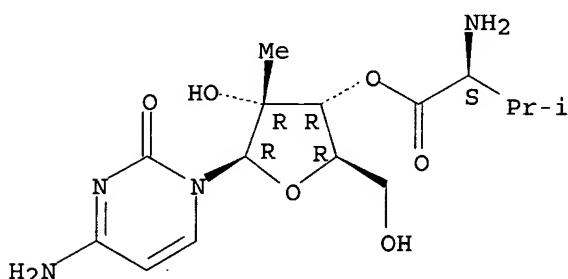
IT 640281-90-9P

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(ribofuranosylcytidine methylvaline ester for treatment of flaviviridae infections)

RN 640281-90-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine (9CI) (CA INDEX NAME)

Absolute stereochemistry.



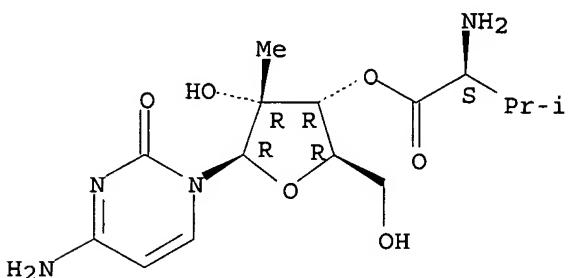
IT 640725-71-9P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(ribofuranosylcytidine methylvaline ester for treatment of flaviviridae infections)

RN 640725-71-9 CAPLUS

CN L-Valine, 3'-ester with 2'-C-methylcytidine, dihydrochloride (9CI) (CA INDEX NAME)

Absolute stereochemistry.



● 2 HCl